**Serial No.:** 09/473,904 **Filed:** December 29, 1999

- i) a substrate with a surface comprising a plurality of assay locations, each assay location comprising an array location comprising a plurality of discrete sites; and
- ii) a population of microspheres comprising at least a first and a second subpopulation each comprising a bioactive agent, wherein said microspheres are distributed on said surface such that said discrete sites each contain no more than one microsphere; and
  - b) determining the presence or absence of said target analyte.
- 19. (Twice Amended) A method of determining the presence of one or more target analytes in one or more samples comprising:
- a) adding said one or more samples to a first substrate comprising a plurality of assay locations, such that said sample is contained at a plurality of said assay locations;
  - b) contacting said sample with a second substrate comprising:
- i) a plurality of array locations, each array location comprising a plurality of discrete sites, wherein at least one assay location is in fluid contact with at least one array location; and
- ii) a population of microspheres comprising at least a first and a second subpopulation each comprising a bioactive agent, wherein said microspheres are distributed on at least one of said array locations such that said discrete sites each contain no more than one microsphere; and
  - c) determining the presence or absence of said target analyte.

(Twice Amended) A method according to claim 18, wherein each of said assay locations comprises a library of bioactive agents.

(Amended) A method according to claim 18, wherein said substrate is a microtiter plate and each assay location is a microtiter well.

(Amended) A method according to claim 18, wherein each discrete site is a bead well.

(Amended) A method according to claim 18, wherein each of said subpopulations further comprise an optical signature capable of identifying said bioactive agent.

**Serial No.:** 09/473,904 **Filed:** December 29, 1999

(Twice Amended) A method according to claim 18, wherein at least a first and second microsphere in said subpopulations further comprise an identifier binding ligand that will bind a decoder binding ligand, whereby said bioactive agent is identified by said identifier binding ligand binding to said decoder binding ligand.

(Amended) A method according to claim 19, wherein said first substrate is a microtiter plate.

26. (Amended) A method according to claim 18 or 28, wherein said second substrate comprises a plurality of fiber optic bundles comprising a plurality of individual fibers, each bundle comprising an array location, and each individual fiber comprising a bead well.

(Amended) A method according to claim 15, wherein each of said subpopulations further comprise an optical signature capable of identifying said bioactive agent.

(Amended) A method according to claim 19, wherein each of said subpopulations further comprise an identifier binding ligand that will bind a decoder binding ligand such that the identification of the bioactive agent can be elucidated.

(Amended) A method according to claim 18 or 19, wherein at least one of said target analytes is a nucleic acid.

Please add Claims 35 and 36:

samples comprising:

- a) contacting said one or more samples with a composition comprising:
- i) a composite array comprising a plurality of assay locations, each assay location comprising an array location comprising a plurality of discrete sites; and
- ii) a population of microspheres comprising at least a first and a second subpopulation each comprising a bioactive agent, wherein said microspheres are distributed on said surface such that said discrete sites each contain no more than one microsphere; and
  - b) determining the presence or absence of said target analyte.



**Serial No.:** 09/473,904 **Filed:** December 29, 1999

- 36. (New) A method of determining the presence of one or more target analytes in one or more samples comprising:
- a) adding said one or more samples to a first substrate comprising a plurality of assay locations, such that said sample is contained at a plurality of said assay locations;
  - b) contacting said sample with a second substrate comprising:
- i) a composite array comprising a plurality of array locations, each array location comprising a plurality of discrete sites, wherein at least one assay location is in fluid contact with at least one array location; and
- ii) a population of microspheres comprising at least a first and a second subpopulation each comprising a bioactive agent, wherein said microspheres are distributed on said surface such that said discrete sites each contain no more than one microsphere; and
  - c) determining the presence or absence of said target analyte.--

